Appln. No. 09/989,388 Amd. dated April 12, 2005 Reply to Office Action of October 20, 2004

Amendments to the Claims

This listing of claims will replace all prior versions, and listings, of claims in the application:

Listing of Claims:

- 1. (Currently Amended) A Lys-Lys binding site I which is a plasminogen fragment consisting of Kringle 1 to Kringle 3 of a human plasminogen with the N-terminal being lysine, which binding site binds to heparin and has the following properties:
 - a. a molecular weight of 38 kDa;
 - b. it is not glycosylated;
- c. it binds to heparin at pH lower than neutral pH but does not bind to heparin at neutral or higher pH, under physiologic ionic conditions;
- d. it inhibits lung tumor metastasis and lung tumor growth but has no ability to inhibit growth of endothelial cells of blood vessels;

wherein said plasminogen fragment is prepared by;

a. preparing Lys-plasminogen from human plasminogen either by adding plasminogen—plasmin to a solution of naturally occurringhuman plasminogen or by incubating naturally occurringhuman plasminogen in the presence of transexamic acid to autolysis;

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- b. treating the Lys-plasminogen obtained in step
 (a) with the elastase to produce fractions of the fragment
 comprising Kringle 1 to Kringle 3;
- c. identifying the fragment of Kringle 1 to Kringle 3 which binds to heparin.
- 2. (Currently Amended) A process for preparing a plasminogen fragment consisting of Kringle 1 to Kringle 3 of a human plasminogen with the N-terminal being lysine, said fragment having the ability to inhibit lung tumor growth, but having no ability to inhibit growth of endothelial cells of blood vessels, comprising;
- a. preparing Lys-plasminogen from naturally occurringhuman plasminogen either by adding plasmin to a solution of naturally occurringhuman plasminogen or by incubating naturally occurringhuman plasminogen in the presence of tranexamic acid to autolysis;
- b. treating the Lys-plasminogen obtained in step
 (a) with elastase to produce fractions of the fragment
 consisting of Kringle 1 to Kringle 3;
- c. identifying the fragment of Kringle 1 to Kringle 3 which binds to heparin; and
 - d. isolating the fragment which binds to heparin.

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- 3. (Previously Presented) The process according to claim 2 wherein the fragment which binds to heparin is recovered by passing a solution of a Lys-plasminogen lysate with elastase through a carrier to which heparin is coupled as a ligand to adsorb those fragments which bind to heparin, and eluting those fragments which do not bind to heparin.
- 4. (Previously Presented) A composition for inhibiting lung tumor metastasis and lung tumor growth comprising an effective amount of a fragment according to claim 1 and, optionally, a pharmaceutically acceptable carrier.

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